## **CLAIMS:**

- A method for inducing melanogenesis in a human subject having an MC1R variant allele associated with loss of or diminished receptor function, which comprises the steps of administering to said subject an amount of an α-MSH analogue effective to induce melanogenesis by the melanocytes in the skin or other epidermal tissue of the subject.
- 2. The method of claim 1, wherein the  $\alpha$ -MSH analogue is selected from:
  - (a) compounds of the formula:

    Ac-Ser-Tyr-Ser-M-Gln-His-D-Phe-Arg-Trp-Gly-Lys-Pro-Val-NH<sub>2</sub>

    wherein M is Met, Nle or Lys; and
  - (b) compounds of the formula: R<sub>1</sub>-W-X-Y-Z-R<sub>2</sub>

wherein

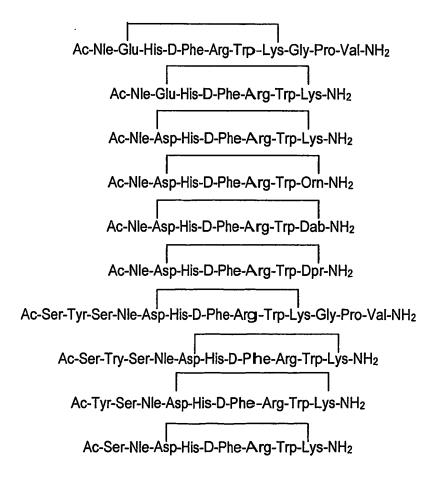
 $R_1$  is Ac-Gly-, Ac-Met-Glu, Ac-Nle-Glu-, or Ac-Tyr-Glu-; W is –His- or –D-His-; X is –Phe-, -D-Phe-, -Tyr-, -D-Tyr-, or -(pNO<sub>2</sub>)D-Phe<sup>7</sup>-; Y is –Arg- or –D-Arg-; Z is –Trp- or –D-Trp-; and  $R_2$  is –NH<sub>2</sub>; -Gly-NH<sub>2</sub>; or –Gly-Lys-NH<sub>2</sub>.

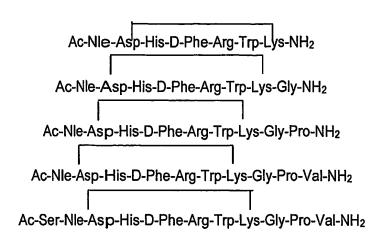
- 3. The method of claim 1, wherein the α-MSH analogue is a cyclic analogue wherein an intramolecular interaction exists (1) between the amino acid residue at position 4 and an amino acid residue at position 10 or 11, and/or (2) between the amino acid residue at position 5 and the amino acid residue at position 10 or 11.
- 4. The method of claim 3, wherein the intramolecular interaction is a disulfide bond or other covalent bond.
- 5. The method of claim 1, wherein the α-MSH analogue is selected from the group consisting of:

Ac-Ser-Tyr-Ser-Nie-Glu-His-D-Phe-Arg-Trp-Lys-Gly-Pro-Val-NH<sub>2</sub>
Ac-Ser-Tyr-Ser-Nie-Asp-His-D-Phe-Arg-Trp-Lys-Gly-Pro-Val-NH<sub>2</sub>

Ac-Nle-Glu-His-D-Phe-Arg-Trp-Lys-Gly-Pro-Val-NH<sub>2</sub>
Ac-Nle-Asp-His-D-Phe-Arg-Trp-Lys-Gly-Pro-Val-NH<sub>2</sub>
Ac-Nle-Asp-His-D-Phe-Arg-Trp-Gly-NH<sub>2</sub>
Ac-Nle-Glu-His-D-Phe-Arg-Trp-Lys-NH<sub>2</sub>
Ac-Nle-Asp-His-D-Phe-Arg-Trp-Orn-NH<sub>2</sub>
Ac-Nle-Asp-His-D-Phe-Arg-Trp-Orn-NH<sub>2</sub>
Ac-Nle-Asp-His-D-Phe-Arg-Trp-Dab-NH<sub>2</sub>
Ac-Nle-Glu-His-D-Phe-Arg-Trp-Dab-NH<sub>2</sub>
Ac-Nle-Asp-His-D-Phe-Arg-Trp-Dab-NH<sub>2</sub>
Ac-Nle-Glu-His-D-Phe-Arg-Trp-Dpr-NH<sub>2</sub>
Ac-Nle-Glu-His-D-Phe-Arg-Trp-Dpr-NH<sub>2</sub>
Ac-Nle-Glu-His-Phe-Arg-Trp-Lys-NH<sub>2</sub>

6. The method of claim 1, wherein the  $\alpha$ -MSH analogue is selected from the group consisting of:





7. The method of claim 1, wherein the  $\alpha$ -MSH analogue is [D-Phe<sup>7</sup>]- $\alpha$ -MSH,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH,

[D-Ser<sup>1</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH,

[D-Tyr<sup>2</sup>, D-Phe<sup>7</sup>]- $\alpha$ --MSH,

[D-Ser<sup>3</sup>, D-Phe<sup>7</sup>]- $\alpha$ --MSH,

[D-Met<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH,

[D-Glu<sup>5</sup>, D-Phe<sup>7</sup>]-α-MSH,

[D-His<sup>6</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH,

[D-Phe<sup>7</sup>, D-Arg<sup>8</sup>]- $\alpha$ -MSH,

[D-Phe<sup>7</sup>, D-Trp<sup>9</sup>]-α-MSH,

[D-Phe<sup>7</sup>, D-Lys<sup>11</sup>]- $\alpha$ -MSH,

[D-Phe-7, D-Pro<sup>12</sup>]- $\alpha$ -MSH,

[D-Phe<sup>7</sup>, D-Val<sup>13</sup>]-α-MSH,

[D-Ser<sup>1</sup>, Nle<sup>4</sup>, D-Phe<sup>7</sup>]-α-MSH,

[D-Tyr<sup>2</sup>, Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH,

[D-Ser<sup>3</sup>, Nle<sup>4</sup>, D-Phe<sup>7</sup>]-α-MSH,

[Nle<sup>4</sup>, D-Glu<sup>5</sup>,D-Phe<sup>7</sup>]-α-MSH,

[Nle<sup>4</sup>, D-His<sup>6</sup>, D-Phe<sup>7</sup>]-α-MSH,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Arg<sup>8</sup>]-α-MSH,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Trp<sup>9</sup>]- $\alpha$ -MSH,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Lys<sup>11</sup>]-α-MSH,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Pro<sup>12</sup>]- $\alpha$ -MSH,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Val<sup>13</sup>]-α-MSH,

[Cys4, Cys10]-α-MSH

[Cys<sup>4</sup>, D-Phe<sup>7</sup>, Cys<sup>10</sup>]-α-MSH

[Cys<sup>4</sup>, Cys<sup>11</sup>]-α-MSH

[Cys<sup>5</sup>, Cys<sup>10</sup>]-α-MSH

[Cys<sup>5</sup>, Cys<sup>11</sup>]-α-MSH

[Cys<sup>4</sup>, Cys<sup>10</sup>]-α-MSH<sub>4-13</sub>

[Cys<sup>4</sup>, Cys<sup>10</sup>]—α-MSH<sub>4-12</sub>

[Nie<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-10</sub>,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>5-11</sub>,

[Nle<sup>4</sup>, D-Tyr<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[(pNO<sub>2</sub>)D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[Tyr<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-10</sub>,

[Tyr<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[NIe4]-α-MSH<sub>4-11</sub>,

[Nle<sup>4</sup>, (pNO<sub>2</sub>)D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[Nle<sup>4</sup>, D-His<sup>6</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[Nle<sup>4</sup>, D-His<sup>6</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[Nle<sup>4</sup>, D-Arg<sup>8</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[Nle<sup>4</sup>, D-Trp<sup>9</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

WO 2005/048967 PCT/AU2004/001630

- 27 -

[Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Trp<sup>9</sup>]- $\alpha$ -MSH<sub>4-11</sub>, [Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-9</sub>, or [Nle<sup>4</sup>, D-Phe<sup>7</sup>, D-Trp<sup>9</sup>]- $\alpha$ -MSH<sub>4-9</sub>.

8. The method of claim 1, wherein the  $\alpha$ -MSH analogue is

[Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-10</sub>,

[Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-11</sub>,

[NIe<sup>4</sup>, D-Phe<sup>7</sup>, D-Trp<sup>9</sup>]- $\alpha$ -MSH<sub>4-11</sub>, or

[Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH<sub>4-9</sub>.

- 9. The method of claim 1, wherein the  $\alpha$ -MSH analogue is [Nle<sup>4</sup>, D-Phe<sup>7</sup>]- $\alpha$ -MSH.
- 10. Use of an α-MSH analogue in the manufacture of a preparation for inducing melanogenesis in a human subject having an MC1R variant allele associated with loss of or diminished receptor function.